DATE: November 13, 1973

To: Dr. Joshua Lederberg

FROM: L. L. Cavalli-Sforza

Subject: Chronology of research on phylogenetic analysis

1961	Work with Anthony Edwards in Pavia begun.
	Three methodsall approximatefor analysis in the case of
	gene frequencies or anthropometric traits (least squares,
	minimum path, cluster analysis. All use matrix of distances
	between population pairs).
1963	Paper at the Hague Genetics Congress.
1964	Anthony and I start on maximum likelihood solution of problems.
196 5	Fitch reinvents independently method 1.
1966	Camin and Sokal independently find approximate method for qualitative
	traits and accept principle that phylogenetic analysis differs from
	standard taxometry.
1966	J. Felsenstein proves covariances between populations are proportional
(unpub1.)	to evolutionary time. Starts on maximum likelihood solution.
1970	Anthony proves that full maximum likelihood solution is practically
	impossible. Gives paper at Royal Statistical Society of London,
	all big brass of British statistics attending. No useful advice
	obtained.
1973	J. Felsenstein publishes iterative maximum likelihood solution
	using matrix of distances.
1973	September, I get interested in measuring "treeness", that is how
	well data are represented by a model of independent evolution. On a
	much law appropriate distribution of innaindependent correlation

problem concerning distribution of non-independent correlation coefficients, I consult with Ingram Olkin. He mentions a formally related problem was solved by a student of Wilks and he himself has provided an extension designed to solve a psychological testing problem. Ingram gives me references to Wilks, Votaw's papers. On this basis, the problem of estimation of tree parameters and of treeness can be solved by the likelihood ratio criterion (a somewhat

treeness can be solved by the likelihood ratio criterion (a somewhat different procedure from classical maximum likelihood but almost as widely accepted). It is basic to use matrix of variances and covariances, not of distances. Solution would have been easy since the beginning if the Wilks' paper—a classic, but rarely read in the original even if it is very readable—had been reinterpreted as an example of a tree with all populations splitting at once.

Some lessons:

- 1) Fundamental papers should be read in the original and not learned from textbooks.
- 2) There is no universal method best for everything.
- One never considers things from sufficiently different angles.